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STRUCTURE FILE UPDATES: 22 AUG 2005 HIGHEST RN 861291-85-2 DICTIONARY FILE UPDATES: 22 AUG 2005 HIGHEST RN 861291-85-2

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 18, 2005

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Structure search iteration limits have been increased. See HELP SLIMITS for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html.

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FILE COVERS 1907 - 23 Aug 2005 VOL 143 ISS 9 FILE LAST UPDATED: 22 Aug 2005 (20050822/ED)

New CAS Information Use Policies, enter HELP USAGETERMS for details.

This file contains CAS Registry Numbers for easy and accurate

substance identification.

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L2
           1481 S L1
              8 S L2 AND ANTIBOD? (S) CONSTRUCT
L3
     ANSWER 1 OF 8 CAPLUS COPYRIGHT 2005 ACS on STN
     Entered STN: 01 Jul 2005
ACCESSION NUMBER:
                         2005:570784 CAPLUS
DOCUMENT NUMBER:
                         143:76835
TITLE:
                         Anti-carcinoembryonic antigen (CEA) single-chain
                         Fv antibodies
                         Fox, Judith A.; Harding, Fiona A.; Schellenberger,
INVENTOR(S):
                         Volker
                         Genencor International, Inc., USA
PATENT ASSIGNEE(S):
SOURCE:
                         PCT Int. Appl., 156 pp.
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
    PATENT NO.
                         KIND
                                DATE
                                           APPLICATION NO.
                                                                   DATE
                                            ______
    WO 2005058236
                         A2
                                20050630
                                         WO 2004-US41429
                                                                   20041210
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA,
             CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI,
             GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP,
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             MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD,
             SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ,
             VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW,
             AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ,
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             NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA,
             GN, GQ, GW, ML, MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.:
                                            US 2003-529354P
                                                                P 20031212
                                            US 2004-577255P
                                                                P 20040604
AΒ
     The present invention relates to anti-(carcinoembryonic antigen)
     single-chain Fv antibodies (CAB mols.), antibody
     -directed enzyme prodrug therapy (ADEPT) constructs directed
     against CEA, and their use in therapy.
IT
     855815-94-0
     RL: PRP (Properties)
        (unclaimed sequence; anti-carcinoembryonic antigen (CEA)
        single-chain Fv antibodies)
    ANSWER 2 OF 8 CAPLUS COPYRIGHT 2005 ACS on STN
L3
    Entered STN: 06 Jan 2005
ACCESSION NUMBER:
                         2005:8670 CAPLUS
DOCUMENT NUMBER:
                         142:87720
                         Myxococcus xanthus genome and proteome sequences
TITLE:
INVENTOR(S):
                         Goldman, Barry S.; Hinkle, Gregory J.; Slater,
                         Steven C.; Wiegand, Roger C.
                         Monsanto Technology, Llc, USA
PATENT ASSIGNEE(S):
SOURCE:
                         U.S., 25 pp.
```

Searcher : Shears 571-272-2528

CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
US 6833447	B1	20041221	US 2001-902540		20010710
US 6833447	B1	20041221	US 2001-902540		20010710
PRIORITY APPLN. INFO.:			US 2000-217883P	P	20000710
			US 2001-902540	А	20010710

AΒ The present invention relates to nucleic acid sequences from the bacterium, Myxococcus xanthus and, in particular, to genomic DNA sequences. Approx. 38,000 genomic nucleotide sequence traces derived from a double-stranded plasmid library prepared from Myxococcus xanthus strain DK1622 are generated and assembled into 1849 contig and singleton sequences, providing a set of about 7842 genes or partial genes and 7134 proteins. A series of predictive and homol. based methods identify proteins involved in polyketide synthesis, serine/threonine protein kinases, antibiotic resistance proteins, DNA modification and restriction enzymes, sigma factors, and nitrate pathway proteins. The invention also encompasses oligonucleotides including primers, e.g. useful for amplifying nucleic acid mols., and collections of nucleic acid mols. and oligonucleotides, e.g. in microarrays. The invention also provides constructs and transgenic cells and organisms comprising nucleic acid mols. of the invention. The invention also relates to methods of using the disclosed nucleic acid mols., oligonucleotides, proteins, fragments of proteins, and antibodies, for example, for gene identification and anal., and preparation of constructs and transgenic cells and organisms. [This abstract record is one of four records for this document necessitated by the large number of index entries required to fully index the document and publication system constraints.].

IT 817294-06-7 817317-86-5

RL: BSU (Biological study, unclassified); BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological study); USES (Uses) (amino acid sequence; Myxococcus xanthus genome and proteome sequences)

ANSWER 3 OF 8 CAPLUS COPYRIGHT 2005 ACS on STN L3

Entered STN: 17 Dec 2004

ACCESSION NUMBER: 2004:1080934 CAPLUS

142:50242 DOCUMENT NUMBER:

Improved expression and secretion of neublastin

using genetic constructs with heterologous signal

peptides and deleting the pro-region

Wahlberg, Lars U.; Groenborg, Mette; Kusk, Philip; INVENTOR(S):

Tornoee, Jens; Pederson, Nels E.; Sisk, William P.

PATENT ASSIGNEE(S): Nsgene A/S, Den.; Biogen Idec Ma, Inc.

SOURCE: PCT Int. Appl., 168 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

TITLE:

PATENT NO. KIND APPLICATION NO. DATE DATE

> Searcher 571-272-2528 : Shears

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WO 2004108760
                                            WO 2004-DK411
                          A2
                                                                    20040610
                                20041216
     WO 2004108760
                         А3
                                20050407
             AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA,
             CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI,
             GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP,
             KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW,
             MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD,
             SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ,
             VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW,
             AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ,
             DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL,
             PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ,
             GW, ML, MR, NE, SN, TD, TG
                                20050428
     US 2005089960
                          Α1
                                            US 2004-864891
     US 2005158824
                          A1
                                20050721
                                            US 2004-957221
                                                                    20041001
PRIORITY APPLN. INFO.:
                                            DK 2003-861
                                                                A 20030610
                                            US 2003-507483P
                                                                   20031002
     The present invention concerns methods and compns. for producing a
AB
     neublastin polypeptide as well as local delivery of neublastin to
     specific regions of the nervous system (including the central nervous
     system and the eye for example) by gene therapy. The biol. active
     neublastin polypeptide is produced from a construct not encoding
     naturally occurring neublastin pro-region, i.e. a construct comprising
     a nucleic acid with a promoter sequence operably linked to a
     nucleotide sequence encoding a signal peptide and a neublastin
     polypeptide, wherein said nucleotide sequence does not encode a
     neublastin pro-region.
IT
     809295-95-2P 809295-96-3P, Neublastin, prepro-
     (human) 809295-99-6P, 25-140-Neublastin (human)
     809296-00-2P, 28-140-Neublastin (human) 809296-08-0P
     809296-09-1P 809296-15-9P 809296-20-6P
     809296-21-7P 809296-26-2P 809296-27-3P
     809296-32-0P 809296-33-1P
     RL: BPN (Biosynthetic preparation); BUU (Biological use,
     unclassified); PRP (Properties); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (amino acid sequence; improved expression and secretion of
        neublastin using genetic constructs with heterologous signal
        peptides and deleting pro-region)
IT
     809297-13-0
     RL: PRP (Properties)
        (unclaimed protein sequence; improved expression and secretion of
        neublastin using genetic constructs with heterologous signal
        peptides and deleting pro-region)
IT
     809297-31-2
     RL: PRP (Properties)
        (unclaimed sequence; improved expression and secretion of
        neublastin using genetic constructs with heterologous signal
        peptides and deleting pro-region)
     ANSWER 4 OF 8 CAPLUS COPYRIGHT 2005 ACS on STN
     Entered STN: 09 Jul 2004
ACCESSION NUMBER:
                         2004:550525 CAPLUS
DOCUMENT NUMBER:
                         141:87790
                         Vector system comprising a nucleotide sequence
TITLE:
```

coding for an antibody

INVENTOR(S): Kingsman, Alan John; Bebbington, Christopher

Robert; Carroll, Miles William; Ellard, Fiona

Margaret; Kingsman, Susan Mary; Myers, Kevin Alan;

Lamikanra, Abigail

PATENT ASSIGNEE(S):

SOURCE:

UK
U.S. Pat. Appl. Publ., 68 pp., Cont.-in-part of

U.S. Pat. Appl. 2003 83,290.

CODEN: USXXCO

DOCUMENT TYPE:

LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT	NO.	KIND	DATE	APPLICATION NO.	DATE
US 2004 WO 2006 WO 2006	0029428	A1 A2 A3	20040708 20000525 20001109	US 2002-334235 WO 1999-GB3859	20021230 19991118.
W:	AE, AL,	AM, AT, AU	, AZ, BA,	BB, BG, BR, BY, CA,	CH, CN, CR,
	CU, CZ,	DE, DK, DM	, EE, ES,	FI, GB, GD, GE, GH,	GM, HR, HU,
				KP, KR, KZ, LC, LK,	
				MW, MX, NO, NZ, PL,	
	SD, SE,	SG, SI, SK	, SL, TJ,	TM, TR, TT, TZ, UA,	UG, US, UZ,
	VN, YU,	ZA, ZW, AM	, AZ, BY,	KG, KZ, MD, RU, TJ,	TM
RW:	GH, GM,	KE, LS, MW	, SD, SL,	SZ, TZ, UG, ZW, AT,	BE, CH, CY,
	DE, DK,	ES, FI, FR	, GB, GR,	IE, IT, LU, MC, NL,	PT, SE, BF,
	BJ, CF,	CG, CI, CM	, GA, GN,	GW, ML, MR, NE, SN,	TD, TG
US 6852	2703	B1	20050208	US 2000-445375	20000321
WO 2001	L036486	A2	20010525	WO 2000-GB4317	20001113
WO 200	L036486	A3	20020510		
w:	AE, AG,	AL, AM, AT	, AU, AZ,	BA, BB, BG, BR, BY,	BZ, CA, CH,
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	GM, HR,	HU, ID, IL	, IN, IS,	JP, KE, KG, KP, KR,	KZ, LC, LK,
	LR, LS,	LT, LU, LV	, MA, MD,	MG, MK, MN, MW, MX,	MZ, NO, NZ,
	PL, PT,	RO, RU, SD	, SE, SG,	SI, SK, SL, TJ, TM,	TR, TT, TZ,
	UA, UG,	US, UZ, VN	, YU, ZA,	ZW, AM, AZ, BY, KG,	KZ, MD, RU,
	TJ, TM				
RW:	GH, GM,	KE, LS, MW	, MZ, SD,	SL, SZ, TZ, UG, ZW,	AT, BE, CH,
	CY, DE,	DK, ES, FI	, FR, GB,	GR, IE, IT, LU, MC,	NL, PT, SE,
	TR, BF,	BJ, CF, CG	, CI, CM,	GA, GN, GW, ML, MR,	NE, SN, TD, TG
US 2003	3083290	A1	20030501	US 2002-60585	20020129
PRIORITY API	PLN. INFO.	:		GB 1997-11579	A 19970604.
				GB 1997-13150	A 19970620
				GB 1997-14230	A 19970704
				WO 1999-GB3859	A 19991118
				GB 2000-3527	A 20000215
				GB 2000-5071	A 20000302
				US 2000-445375	A2 20000321
				WO 2000-GB4317	A2 20001113
				US 2002-60585	A2 20020129

WO 1998-GB1627	W	19980604
GB 1998-25303	Α	19981118
GB 1999-1739	Α	19990127
GB 1999-17995	Α	19990730

AB The authors disclose an expression vector cassette system comprising a nucleotide sequence coding for an antibody. In one example, the nucleotide sequence encodes a single-chain construct of an antibody targeted to the oncofetal glycoprotein 5T4.

IT 149298-29-3

RL: PRP (Properties)

(unclaimed sequence; vector system comprising a nucleotide sequence coding for an antibody)

L3 ANSWER 5 OF 8 CAPLUS COPYRIGHT 2005 ACS on STN

ED Entered STN: 24 Oct 2003

ACCESSION NUMBER: 2003:837306 CAPLUS

DOCUMENT NUMBER:

139:334817

TITLE:

Identifying modulators for serine/threonine kinases using phospho-specific antibodies and kinase proteins fused with substrate proteins

INVENTOR(S):

Suda, Mikiya; Shibahara, Megumi

PATENT ASSIGNEE(S):

Glaxo Group Limited, UK PCT Int. Appl., 38 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

SOURCE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.					KIND DATE			APPLICATION NO.							DATE		
W	70 21	2003087394			A1 2003102			1023	. 1	WO 2	003-1		20030415				
	1	<i>N</i> :	ΑE,	ΑG,	AL,	AM,	ΑT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,
			CN,	co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,
			GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	ΚP,	KR,	KZ,
			LC,	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,
			NO,	ΝZ,	OM,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	TJ,
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]	RW:	GH,	GM,	ΚE,	LS,	MW,	ΜZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,
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			EE,	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,
			SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	G₩,	ML,	MR,
			ΝE,	SN,	TD,	TG											•
PRIORI					US 2002-372662P						P 20020415						

AB A method is provided for identifying a modulator for serine/threonine kinase employing expression of fusion proteins of a kinase substrate, such as p53, and the serine/threonine kinase. The fusion protein between a substrate protein and a serine/threonine kinase is expressed is a cell, the cell incubated with a candidate modulator, and the level of phosphorylation of the substrate determined Phosphorylation may be determined in may ways, including Western blotting and ELISA using phospho-specific antibodies. Protein p53 is chosen as substrate within the fusion protein construct based on the fact that

(1) p53 can be phosphorylated by multiple kinases such as Chk1/2, protein kinase A, and JNK kinase; (2) within p53 there are several phosphorylation sites; and (3) antibodies are com. available which recognize a particular phosphorylation site within p53. Preferred kinases can be derived from p38, JNK3, SGK, PLK1, YAK3, MAPKAPK2, MYT1, CDK5, ROCK1/2, and Chk1.

IT 615863-97-3 616522-70-4 616522-72-6 616522-74-8

RL: ANT (Analyte); ARG (Analytical reagent use); PRP (Properties); ANST (Analytical study); USES (Uses)

(amino acid sequence; identifying modulators for serine/threonine kinases using phospho-specific antibodies and kinase proteins fused with substrate proteins)

REFERENCE COUNT:

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 6 OF 8 CAPLUS COPYRIGHT 2005 ACS on STN

8

ED Entered STN: 20 Mar 2003

ACCESSION NUMBER: 2003:219693 CAPLUS

DOCUMENT NUMBER: 138:253715

TITLE: Multimeric single chain tandem Fv-antibodies

INVENTOR(S): Le Gall, Fabrice; Kipriyanov, Sergey; Reusch, Uwe;

Moldenhauer, Gerhard; Little, Melvyn

PATENT ASSIGNEE(S): Affimed Therapeutics AG, Germany

SOURCE:

Eur. Pat. Appl., 29 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.					KIND DATE		APPLICATION NO.							DATE				
E	P.	1293	514			A1	_	2003	0319		EP 2	001-	1221	04		2	0010	914
		R:	•	•	•			ES,	•	-	-		-	LU,	NL,	SE,	MC,	
T-0		2002		-				FI,						207		2	กกวก	012
										WO 2002-EP10307							0020	913
W	10	2003	0250:	18		A 3		2003	0828									
		W:	ΑE,	AG,	AL,	AM,	AT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	
			CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	
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			NO,	NZ,	OM,	PH,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	ТJ,	
			TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VN,	YU,	ZA,	ZM,	ZW		
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			EE,	ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	SK,	TR,	
			BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG
J	JΡ	2005	5086	28		Т2		2005	0407		JP 2	003-	5288	63		2	0020	913
U	IS	2005	0791	70		A 1		2005	0414		US 2	003-	4896	26		2	0020	913
PRIORI	TY	APP	LN.	INFO	.:						EP 2	001-	1221	04	j	A 2	0010	914
											WO 2	002-	EP10	307	1	w 2	0020	913

AB The authors disclose the preparation and characterization of multimeric tandem scFv-antibody constructs. A monomeric construct is comprised of one single-chain Fv fragment joined by a linker peptide to a second single-chain Fv fragment of the same or

different specificity. The linker peptides joining each pair of VH/VL domains in the tandem construct are not identical; the monomer contains a terminal dimerization domain to facilitate multimerization. In one example, a tandem scFv-antibody construct was prepared with specificity for CD3 and CD19.

IT 502771-42-8P 502771-44-0P

> RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified); DGN (Diagnostic use); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(amino acid sequence; preparation and characterization of multimeric tandem single-chain antibodies)

IT 149298-29-3

RL: PRP (Properties)

(linker peptide for preparation of multimeric tandem single-chain antibodies)

REFERENCE COUNT:

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 7 OF 8 CAPLUS COPYRIGHT 2005 ACS on STN L3

Entered STN: 08 Sep 2002 ED

ACCESSION NUMBER:

2002:676155 CAPLUS

DOCUMENT NUMBER:

137:214226

TITLE:

Use of mammalian retinoid-inducible serine

carboxypeptidase gene in diagnosis and treatment

of vascular diseases

INVENTOR(S):

Miano, Joseph Michael; Streb, Jeffrey Williams;

Chen, Jiyuan

PATENT ASSIGNEE(S):

University of Rochester, USA

SOURCE:

PCT Int. Appl., 129 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PATENT NO.					KIND DATE			APPLICATION NO.							DATE	
WO	2002	0685	99			20020906			1				2	0020222			
WO							AU,		ת כו	DD	BC.	DD	ΒV	D 7	רא	CH	
	w:	•	•	•	•		•	•	•	•	•	•	-	-			
		•	•	•	•		DE,	•		•		-	-		-	•	
	•	•	•	•	•		ID,	•	•	•	•	•	•	-		•	
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		NO,	ΝZ,	OM,	PH,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	TJ,	
		TM,	TN,	TR,	TT,	TZ,	UA,	ŪG,	US,	UZ,	VN,	ΥU,	ZA,	ZM,	ZW		
	RW:	GH,	GM,	ΚE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	ŪG,	ZM,	ZW,	AM,	AZ,	
		BY,	KG,	KZ,	MD,	RU,	TJ,	TM,	AT,	BE,	CH,	CY,	DE,	DK,	ES,	FI,	
		FR,	GB,	GR,	IE,	IT,	LU,	MC,	NL,	PT,	SE,	TR,	BF,	ВJ,	CF,	CG,	
							GW,										
CA	2438	827 [°]	•	-	AA		2002	0906	•	CA 2	002-	2438	827		2	0020222	
																0020222	
							ES,										
					CY,		•	•	•	•							
л US	2004						2004	1007		US 2	004-	4686	55		2	0040423	
PRIORIT																0010222	
										บร 2	001-	2930	97P		P 2	0010523	

WO 2002-US5560 W 20020222

The present invention relates to an isolated retinoid inducible serine carboxypeptidase proteins or polypeptides, and the nucleic acid mols. encoding such a protein or polypeptide. Nucleic acid constructs, expression systems and host cells containing those nucleic acid mols., and antibodies raised against the proteins or polypeptides are also disclosed. The present invention also relates to methods for detecting a vascular disease or disorder, inhibiting smooth muscle cell growth, treating vascular hyperplasia, and inhibiting the activity of extracellular regulated kinase. The present invention also relates to a transgenic non-human animal lacking a gene encoding a retinoid inducible protein or polypeptide.

IT 456543-55-8 457048-38-3

RL: PRP (Properties)

(unclaimed sequence; mammalian retinoid-inducible serine carboxypeptidase gene in diagnosis and treatment of vascular diseases)

L3 ANSWER 8 OF 8 CAPLUS COPYRIGHT 2005 ACS on STN

ED Entered STN: 12 Nov 1999

ACCESSION NUMBER: 1999:723065 CAPLUS

DOCUMENT NUMBER: 131:350246

TITLE:

Multivalent Fv antibody

constructs containing at least four

variable domains and their use in diagnosis and

therapy

INVENTOR(S):

Little, Melvyn; Kipriyanov, Sergej

PATENT ASSIGNEE(S):

Deutsches Krebsforschungszentrum Stiftung des

Offentlichen Rechts, Germany

SOURCE:

PCT Int. Appl., 48 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.					KIND DATE			į	APPL	ICAT:		DATE				
	9957								1	WO 1	999-1		19990505			
WO	9957															
	W:	AL,	AM,	ΑT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CU,	CZ,
		DK,	EE,	ES,	FI,	GB,	GD,	GΕ,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,
		JP,	ΚE,	KG,	KΡ,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	MD,	MG,
		MK,	MN,	MW,	MX,	NO,	NZ,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,
		SL,	TJ,	TM,	TR,	TT,	UA,	UG,	US,	UZ,	VN,	YU,	ZW,	AM,	AZ,	BY,
					RU,			·	•	•	•	٠.	•			
	RW:	•		•	•	•	SD,	SL,	SZ.	ŬĠ,	ZW,	AT,	BE,	CH,	CY,	DE,
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WO 1999-DE1350 W 19990505

AB The invention relates to a multivalent Fv antibody construct comprising at least four variable domains which are connected to one another via peptide linkers 1, 2 and 3. The invention also relates to expression plasmids which code for such an Fv antibody construct. In addition, the invention relates to a method for producing the Fv antibody constructs and to the use thereof. The invention is based on the discovery that the stability of Fv constructs are enhanced when they are in the form of single-chain dimers in which the 4 variable domains are connected via 3 peptide linkers. The Fv construct folds with itself when the middle peptide linker contains 10-30 amino acids. However, when the middle peptide linker contains 10 or fewer amino acids the Fv construct folds with another Fv construct thus producing a multivalent multimer. These Fv constructs may addnl. be multispecific. For example, Fv dimers and tetramers targeting both CD3 and CD19 antigens were prepared with recombinant Escherichia coli and Pichia pastoris.

IT 250245-29-5P 250245-35-3P

RL: ARG (Analytical reagent use); BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); PREP (Preparation); USES (Uses)

(amino acid sequence; multivalent Fv antibody constructs containing at least four variable domains and their

IT 149298-29-3 249509-02-2

use in diagnosis and therapy)

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence)

(peptide linker; multivalent Fv antibody constructs containing at least four variable domains and their use in diagnosis and therapy)

E31 THROUGH E60 ASSIGNED

FILE 'REGISTRY' ENTERED AT 14:46:27 ON 23 AUG 2005

L4

30 SEA FILE=REGISTRY ABB=ON PLU=ON (149298-29-3/BI OR 249509-02-2/BI OR 250245-29-5/BI OR 250245-35-3/BI OR 456543-55-8/BI OR 457048-38-3/BI OR 502771-42-8/BI OR 502771-44-0/BI OR 615863-97-3/BI OR 616522-70-4/BI OR 616522-72-6/BI OR 616522-74-8/BI OR 809295-95-2/BI OR 809295-96-3/BI OR 809295-99-6/BI OR 809296-00-2/BI OR 809296-08-0/BI OR 809296-09-1/BI OR 809296-15-9/BI OR 809296-20-6/BI OR 809296-21-7/BI OR 809296-26-2/BI OR 809296-27-3/BI OR 809296-32-0/BI OR 809296-33-1/BI OR 809297-13-0/BI OR 809297-31-2/BI OR 817294-06-7/BI OR 817317-86-5/BI OR 855815-94-0/BI)

- L4 ANSWER 1 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN
- RN 855815-94-0 REGISTRY
- CN 59: PN: WO2005058236 PAGE: 62/79 unclaimed protein (9CI) (CA INDEX NAME)
- SOL 622
- MF Unspecified
- CI MAN

REFERENCE 1: 143:76835

L4 ANSWER 2 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN

817317-86-5 REGISTRY RN CN Protein MYX12 4863 (Myxococcus xanthus strain DK1622) (9CI) (CA INDEX OTHER NAMES: 2553: PN: US6833447 SEQID: 14554 claimed protein CN SQL MF Unspecified MAN CI REFERENCE 1: 142:87720 ANSWER 3 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4817294-06-7 REGISTRY RNProtein MYX12 2448 (Myxococcus xanthus strain DK1622) (9CI) (CA INDEX CN NAME) OTHER NAMES: 139: PN: US6833447 SEQID: 12139 claimed protein CN SQL 198 MF Unspecified CI MAN REFERENCE 1: 142:87720 ANSWER 4 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4RN809297-31-2 REGISTRY 84: PN: WO2004108760 TABLE: 1 unclaimed sequence (9CI) (CA INDEX CN NAME) SQL 220 Unspecified MF CI MAN REFERENCE 1: 142:50242 ANSWER 5 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4RN **809297-13-0** REGISTRY 21: PN: WO2004108760 SEQID: 21 unclaimed protein (9CI) CN (CA INDEX NAME) SQL 140 ΜF Unspecified CI MAN REFERENCE 1: 142:50242 ANSWER 6 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4RN **809296-33-1** REGISTRY Somatotropin (human signal peptide) fusion protein with CN 28-140-neublastin (human) (9CI) (CA INDEX NAME) OTHER NAMES: 53: PN: WO2004108760 SEQID: 53 claimed protein CN SQL 139 MF Unspecified CI MAN

REFERENCE 1: 142:50242

ANSWER 7 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4

RN 809296-32-0 REGISTRY

Somatotropin (human signal peptide) fusion protein with neublastin CN (human) (9CI) (CA INDEX NAME)

> Searcher Shears 571-272-2528 :

OTHER NAMES: 52: PN: WO2004108760 SEQID: 52 claimed protein SQL 166 MF Unspecified CI MAN REFERENCE 1: 142:50242 ANSWER 8 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4**809296-27-3** REGISTRY RN CN Signal peptide (synthetic) fusion protein with 28-140-neublastin (human) (9CI) (CA INDEX NAME) OTHER NAMES: 46: PN: WO2004108760 SEQID: 46 claimed protein CN SQL 132 MF Unspecified CI MAN REFERENCE 1: 142:50242 L4ANSWER 9 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN RN809296-26-2 REGISTRY CN Signal peptide (synthetic) fusion protein with neublastin (human) (9CI) (CA INDEX NAME) OTHER NAMES: 45: PN: WO2004108760 SEQID: 45 claimed protein CN SQL 159 ΜF Unspecified CI MAN REFERENCE 1: 142:50242 L4ANSWER 10 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN 809296-21-7 REGISTRY RN Albumin (Rattus norvegicus signal peptide) fusion protein with CN 28-140-neublastin (human) (9CI) (CA INDEX NAME) OTHER NAMES: CN 39: PN: WO2004108760 SEQID: 39 claimed protein SQL 131 MFUnspecified MAN CI REFERENCE 1: 142:50242 L4ANSWER 11 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN RN 809296-20-6 REGISTRY CN Albumin (Rattus norvegicus signal peptide) fusion protein with neublastin (human) (9CI) (CA INDEX NAME) OTHER NAMES: 38: PN: WO2004108760 SEQID: 38 claimed protein CN SQL 158 ΜF Unspecified MAN CI REFERENCE 1: 142:50242 ANSWER 12 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4 RN**809296-15-9** REGISTRY CN Immunoglobulin (Mus musculus signal peptide) fusion protein with

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Shears

571-272-2528

neublastin (human) (9CI) (CA INDEX NAME) OTHER NAMES: 31: PN: W02004108760 SEQID: 31 claimed protein SOL Unspecified MF MAN CI REFERENCE 1: 142:50242 ANSWER 13 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4809296-09-1 REGISTRY RN Neublastin (human signal peptide) fusion protein with CN 28-140-neublastin (human) (9CI) (CA INDEX NAME) OTHER NAMES: 26: PN: WO2004108760 SEQID: 26 claimed protein CN SQL 152 MF Unspecified CI MAN REFERENCE 1: 142:50242 ANSWER 14 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4809296-08-0 REGISTRY RN Neublastin (human signal peptide) fusion protein with neublastin (human) (9CI) (CA INDEX NAME) OTHER NAMES: (1-39), (81-220) - Neublastin, prepro- (human) CN CN 25: PN: WO2004108760 SEQID: 25 claimed protein SQL 179 MF Unspecified MAN CI REFERENCE 1: 142:50242 ANSWER 15 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4809296-00-2 REGISTRY RN 28-140-Neublastin (human) (9CI) (CA INDEX NAME) OTHER NAMES: 14: PN: WO2004108760 SEQID: 14 claimed protein CN SOL 113 MF Unspecified CI MAN REFERENCE 1: 142:50242 ANSWER 16 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4RN 809295-99-6 REGISTRY CN 25-140-Neublastin (human) (9CI) (CA INDEX NAME) OTHER NAMES: 13: PN: WO2004108760 SEQID: 13 claimed protein CNSQL MF Unspecified CI MAN REFERENCE 1: 142:50242 ANSWER 17 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4 809295-96-3 REGISTRY RN Neublastin, prepro- (human) (9CI) (CA INDEX NAME) CN

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571-272-2528

OTHER NAMES: 10: PN: WO2004108760 SEQID: 10 claimed protein SQL 220 MF Unspecified MAN CI REFERENCE 1: 142:50242 ANSWER 18 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4809295-95-2 REGISTRY RNImmunoglobulin (Mus musculus immunoglobulin) fusion protein with CN 28-140-neublastin (human) (9CI) (CA INDEX NAME) OTHER NAMES: 9: PN: WO2004108760 SEOID: 9 claimed protein CN SQL 132 MF Unspecified CI MAN REFERENCE 1: 142:50242 ANSWER 19 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4616522-74-8 REGISTRY RNp53 (protein) (human) fusion protein with 1-270-gene chk1 protein CN kinase (human) (9CI) (CA INDEX NAME) OTHER NAMES: 7: PN: WO03087394 SEQID: 7 claimed protein CN SQL 686 MF Unspecified CI MAN REFERENCE 1: 139:334817 ANSWER 20 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4RN616522-72-6 REGISTRY p53 (protein) (human) fusion protein with 60-431-gene sgk protein kinase (human) (9CI) (CA INDEX NAME) OTHER NAMES: 5: PN: WOO3087394 SEQID: 5 claimed protein CN SQL 788 MF Unspecified CI MAN REFERENCE 1: 139:334817 ANSWER 21 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4616522-70-4 REGISTRY RN p53 (protein) (human) fusion protein with 1-378-gene Myt1 protein CN kinase (human) (9CI) (CA INDEX NAME) OTHER NAMES: 3: PN: WO03087394 SEQID: 3 claimed protein CN SOL 794 MF Unspecified CI MAN REFERENCE 1: 139:334817 ANSWER 22 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN L4615863-97-3 REGISTRY RN

Searcher: Shears 571-272-2528

p53 (protein) (human) fusion protein with gene YAK3 protein kinase

CN

```
(human) (9CI) (CA INDEX NAME)
OTHER NAMES:
     1: PN: WO03087394 SEQID: 1 claimed protein
CN
SQL
MF
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CI
     MAN
REFERENCE
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     ANSWER 23 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN
L4
RN
     502771-44-0 REGISTRY
     Immunoglobulin, anti-(human CD19 (antigen)) (mouse clone DSM-14471
CN
     single-chain precursor) fusion protein with immunoglobulin,
     anti-(human CD3 (antigen)) (mouse single-chain) (9CI) (CA INDEX NAME)
OTHER NAMES:
     3: PN: EP1293514 FIGURE: 7 claimed protein
SQL 562
     Unspecified
MF
CI
     MAN
REFERENCE
            1: 138:253715
T.4
     ANSWER 24 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN
RN
     502771-42-8 REGISTRY
CN
     Immunoglobulin, anti-(human CD3 (antigen)) (mouse clone DSM-14470
     single-chain precursor) fusion protein with immunoglobulin,
     anti-(human CD19 (antigen)) (mouse single-chain) (9CI) (CA INDEX
     NAME)
OTHER NAMES:
     1: PN: EP1293514 FIGURE: 6 claimed protein
SQL 562
     Unspecified
MF
     MAN
CI
REFERENCE
            1: 138:253715
     ANSWER 25 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN
L4
     457048-38-3 REGISTRY
RN
     L-Serine, L-tryptophyl-L-leucyl-L-glutaminylglycylglycyl-L-
CN
     prolylglycyl-L-seryl-L-seryl- (9CI) (CA INDEX NAME)
OTHER NAMES:
     2: PN: WO02068599 SEQID: 2 unclaimed sequence
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     ANSWER 26 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN
L4
RN
     456543-55-8 REGISTRY
CN
     L-Proline, L-tryptophyl-L-leucyl-L-glutaminylglycylglycyl-L-
     prolylglycyl-L-seryl-L-seryl-L-isoleucyl-L-phenylalanyl-L-
     cysteinyl-L-\arglutamyl-L-seryl-L-tyrosylglycylglycyl-L-valyl-L-
     tyrosyl-L-asparaginylglycyl-L-asparaginyl-L-\alpha-glutamyl-L-\alpha-
     aspartyl-L-leucyl-L-isoleucyl-L-leucyl-L-glutaminyl-L-phenylalanyl-L-
     tryptophyl-L-tryptophyl-L-isoleucyl-L-leucyl-L-arginyl-L-alanylglycyl-
     L-histidyl-L-methionyl-L-valyl-L-alanyl-L-tyrosyl-L-α-aspartyl-L-
     threonyl- (9CI) (CA INDEX NAME)
OTHER NAMES:
     74: PN: WO02068599 FIGURE: 1 unclaimed sequence
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45
SQL
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CI
     MAN
REFERENCE
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     ANSWER 27 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN
T.4
RN
     250245-35-3 REGISTRY
CN
     Immunoglobulin, anti-(human CD3 antigen/CD19 antigen) (mouse clone
     pDISC3x19-SL Fv fragment tetramer precursor) fusion protein with gene
     c-myc protein epitope fusion protein with hexahistidine peptide (9CI)
     (CA INDEX NAME)
OTHER NAMES:
     4: PN: WO9957150 SEQID: 4 claimed protein
SQL 539
MF
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CI
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            1: 131:350246
REFERENCE
     ANSWER 28 OF 30 REGISTRY COPYRIGHT 2005 ACS on STN
T.4
     250245-29-5 REGISTRY
RN
     Immunoglobulin, anti-(human CD3 antigen/CD19 antigen) (mouse clone
CN
     pDISC3x19-LL Fv fragment dimer precursor) fusion protein with gene
     c-myc protein epitope fusion protein with hexahistidine peptide (9CI)
     (CA INDEX NAME)
OTHER NAMES:
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     2: PN: WO9957150 SEQID: 2 claimed protein
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     L-Serine, glycylglycyl-L-prolylglycyl- (9CI) (CA INDEX NAME)
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OTHER NAMES:
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     116: PN: US6297041 SEQID: 116 unclaimed protein
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     118: PN: WO0024913 SEQID: 116 claimed protein
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     121: PN: US20040002450 SEQID: 123 unclaimed protein
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     123: PN: US20040001822 SEQID: 123 unclaimed protein
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     125: PN: US20040001839 SEQID: 123 unclaimed protein
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Searcher : Shears 571-272-2528

149: PN: US20050112642 SEQID: 149 unclaimed sequence 151: PN: US20050009750 SEQID: 149 unclaimed sequence

159: PN: WO02053700 SEQID: 123 unclaimed protein

CN

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FILE 'MEDLINE' ENTERED AT 14:47:08 ON 23 AUG 2005
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FILE 'HOME' ENTERED AT 14:47:13 ON 23 AUG 2005

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FILE 'REGISTRY' ENTERED AT 14:44:07 ON 23 AUG 2005
L1 3715 SEA ABB=ON PLU=ON GGGGSGGGGGGGGGGGGGGGGGGGGS/SQSP

FILE 'CAPLUS' ENTERED AT 14:44:40 ON 23 AUG 2005

L2 1481 SEA ABB=ON PLU=ON L1
L*** DEL 5 S L2 AND LITTLE ?/AU
L*** DEL 3 S L3 AND ANTIBOD?
D TI AU 1-3

L*** DEL 530 S L2 AND ANTIBOD? L*** DEL 30 S L3 AND CONSTRUCT

L3 8 SEA ABB=ON PLU=ON L2 AND ANTIBOD? (S) CONSTRUCT
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SEL HIT L3 1-8 RN

FILE 'REGISTRY' ENTERED AT 14:46:27 ON 23 AUG 2005

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30 SEA ABB=ON PLU=ON (149298-29-3/BI OR 249509-02-2/BI OR 250245-29-5/BI OR 250245-35-3/BI OR 456543-55-8/BI OR 457048-38-3/BI OR 502771-42-8/BI OR 502771-44-0/BI OR 615863-97-3/BI OR 616522-70-4/BI OR 616522-72-6/BI OR 616522-74-8/BI OR 809295-95-2/BI OR 809295-96-3/BI OR 809295-99-6/BI OR 809296-00-2/BI OR 809296-08-0/BI OR 809296-09-1/BI OR 809296-15-9/BI OR 809296-20-6/BI OR 809296-21-7/BI OR 809296-26-2/BI OR 809296-27-3/BI OR 809296-32-0/BI OR 809296-33-1/BI OR 809297-13-0/BI OR 809297-31-2/BI OR 817294-06-7/BI OR 817317-86-5/BI OR 855815-94-0/BI)

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FILE 'HOME' ENTERED AT 14:47:13 ON 23 AUG 2005

FILE REGISTRY

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 22 AUG 2005 HIGHEST RN 861291-85-2 DICTIONARY FILE UPDATES: 22 AUG 2005 HIGHEST RN 861291-85-2

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 18, 2005

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- * The CA roles and document type information have been removed from *
- * the IDE default display format and the ED field has been added,

FILE RELOADED: 19 October 2003.

FILE EMBASE

FILE COVERS 1974 TO 18 Aug 2005 (20050818/ED)

EMBASE has been reloaded. Enter HELP RLOAD for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

FILE HOME

* effective March 20, 2005. A new display format, IDERL, is now

 * available and contains the CA role and document type information. *

Structure search iteration limits have been increased. See HELP SLIMIfor details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

FILE CAPLUS

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FILE COVERS 1907 - 23 Aug 2005 VOL 143 ISS 9 FILE LAST UPDATED: 22 Aug 2005 (20050822/ED)

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This file contains CAS Registry Numbers for easy and accurate substance identification.

FILE MEDLINE

FILE LAST UPDATED: 20 AUG 2005 (20050820/UP). FILE COVERS 1950 TO DA

On December 19, 2004, the 2005 MeSH terms were loaded.

The MEDLINE reload for 2005 is now available. For details enter HELP RLOAD at an arrow promt (=>). See also:

http://www.nlm.nih.gov/mesh/ http://www.nlm.nih.gov/pubs/techbull/nd04/nd04 mesh.html

OLDMEDLINE now back to 1950.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2005 vocabulary.

This file contains CAS Registry Numbers for easy and accurate substance identification.

FILE BIOSIS

FILE COVERS 1969 TO DATE.

CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 17 August 2005 (20050817/ED)